



10 / 522609
PCT/GB 2003 / 003206

REC'D 15 AUG 2003	
WIPO	PCT



INVESTOR IN PEOPLE

The Patent Office
Concept House
Cardiff Road 25 JAN 2005
Newport
South Wales
NP10 8QQ

PRIORITY DOCUMENT

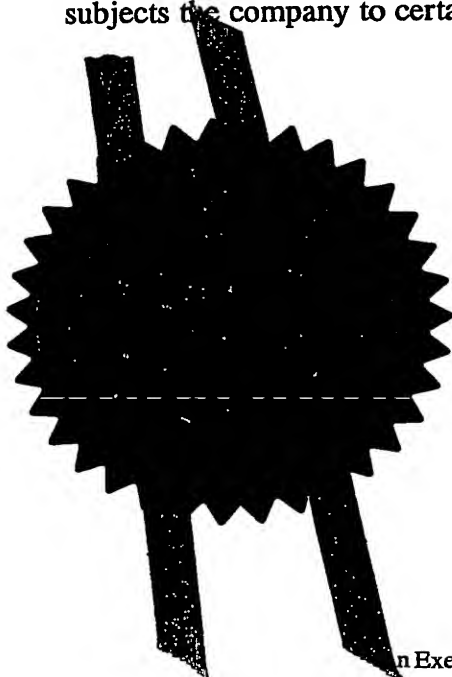
SUBMITTED OR TRANSMITTED IN
COMPLIANCE WITH RULE 17.1(a) OR (b)

I, the undersigned, being an officer duly authorised in accordance with Section 74(1) and (4) of the Deregulation & Contracting Out Act 1994, to sign and issue certificates on behalf of the Comptroller-General, hereby certify that annexed hereto is a true copy of the documents as originally filed in connection with the patent application identified therein.

In accordance with the Patents (Companies Re-registration) Rules 1982, if a company named in this certificate and any accompanying documents has re-registered under the Companies Act 1980 with the same name as that with which it was registered immediately before re-registration save for the substitution as, or inclusion as, the last part of the name of the words "public limited company" or their equivalents in Welsh, references to the name of the company in this certificate and any accompanying documents shall be treated as references to the name with which it is so re-registered.

In accordance with the rules, the words "public limited company" may be replaced by p.l.c., plc, P.L.C. or PLC.

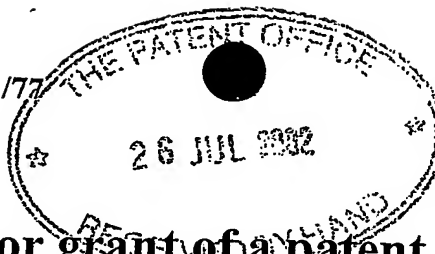
Re-registration under the Companies Act does not constitute a new legal entity but merely subjects the company to certain additional company law rules.



Signed

Dated 8 August 2003

BEST AVAILABLE COPY



The
Patent
Office

1/77

29JUL02 E736529-1 D01631
P01/7700 0.00-0217390.4

The Patent Office

26 JUL 2002

Cardiff Road
Newport
Gwent NP9 1RH

Fee: £0

Request for grant of a patent

(See the notes on the back of this form. You can also get an explanatory leaflet from the Patent Office to help you fill in this form.)

1. Your reference

43232.GB01/SJNG/NT

2. Patent application number

(The Patent Office will fill in this part)

0217390.4

3. Full name, address and postcode of the or of each applicant (underline all surnames)

LEE, Helen
University of Cambridge
Department of Haematology - Diagnostic
Development
East Anglia Blood Centre Site
Long Road
Cambridge
CB2 2PT

Patents ADP number (if you know it)

If the applicant is a corporate body, give the country/state of incorporation

7936131001

4. Title of the invention

Liquid Sampler and Method

5. Full name, address and postcode in the United Kingdom to which all correspondence relating to this form and translation should be sent

Reddie & Grose
16 Theobalds Road
LONDON
WC1X 8PL

Patents ADP number (if you know it)

91001

6. If you are declaring priority from one or more earlier patent applications, give the country and the date of filing of the or of each of these earlier applications and (if you know it) the or each application number

Country

Priority application
(If you know it)

Date of filing
(day/month/year)

7. If this application is divided or otherwise derived from an earlier UK application, give the number and the filing date of the earlier application

Number of earlier application

Date of filing
(day/month/year)

8. Is a statement of inventorship and of right to grant of a patent required in support of this request? (Answer 'Yes' if:

a) any applicant named in part 3 is not an inventor, or

b) there is an inventor who is not named as an applicant, or

c) any named applicant is a corporate body.
See note (d))

YES

Patents Form 1/77

9. Enter the number of sheets for any of the following items you are filing with this form. Do not count copies of the same document.

Continuation sheets of this form

Description	19
Claim(s)	4
Abstract	1
Drawing(s)	12 d. 2 8

10. If you are also filing any of the following, state how many against each item.

Priority documents

Translations of priority documents

Statement of inventorship and right to grant of a patent (*Patents Form 7/77*)

Request for preliminary examination and search (*Patents Form 9/77*)

Request for substantive examination (*Patents Form 10/77*)

Any other documents (please specify)

11.

I/We request the grant of a patent on the basis of this application.

Signature

Reddie & Grose

Date

11 July 2002

REDDIE & GROSE

12. Name and daytime telephone number of person to contact in the United Kingdom

S J N GOODMAN
01223 360350

Warning

After an application for a patent has been filed, the Comptroller of the Patent Office will consider whether publication or communication of the invention should be prohibited or restricted under Section 22 of the Patents Act 1977. You will be informed if it is necessary to prohibit or restrict your invention in this way. Furthermore, if you live in the United Kingdom, Section 23 of the Patents Act 1977 stops you from applying for a patent abroad without first getting written permission from the Patent Office unless an application has been filed at least 6 weeks beforehand in the United Kingdom for a patent for the same invention and either no direction prohibiting publication or communication has been given, or such direction has been revoked.

Notes

- a) If you need help to fill in this form or you have any questions, please contact the Patent Office on 0645 500505.
- b) Write your answers in capital letters using black ink or you may type them.
- c) If there is not enough space for all the relevant details on any part of this form, please continue on a separate sheet of paper and write "see continuation sheet" in the relevant part(s). Any continuation sheet should be attached to this form.
- d) If you have answered 'Yes' Patents Form 7/77 will need to be filed.
- e) Once you have filled in the form you must remember to sign and date it.
- f) For details of the fee and ways to pay please contact the Patent Office.

LIQUID SAMPLER AND METHOD

This invention relates to a liquid sampler and a method for
collecting a liquid sample, and to a diagnostic method
5 comprising the collection of urine.

It is well known that sampling urine can be an effective
research and diagnosis tool to investigate an individual's
physical condition.

10

Summary of Invention

The invention provides in a first aspect a liquid sampler and
a method for sampling a liquid as defined in the appended
15 independent claims. Preferred or advantageous features of the
invention are set out in dependent sub-claims.

As noted above, it is known to perform diagnostic tests on
urine samples. In a second aspect of the invention, however,
20 the inventor has appreciated that significant advantages can
be achieved and new diagnostic tests implemented if a sample
of only the first portion of the urine provided by an
individual is tested or investigated. The first urine portion
may be termed first-void urine.

25

There is a problem, however, in isolating first-void urine
without excessive dilution by the remainder of the contents of
the patient's bladder.

30 Thus, a preferred embodiment of the first aspect of the-
invention may advantageously provide a sampler for sampling a
first portion of a liquid flow. The liquid flow enters at an
inlet of the sampler. The sample passes from the inlet
through a valve into a sample chamber and then the valve
35 closes, diverting the remainder of the liquid flow to an
overflow or exhaust passage. The valve comprises a valve

inlet coupled to the sampler inlet, a valve outlet opening into the sample chamber and positioned, in use, substantially below the valve inlet, a shut-off chamber separating the valve outlet from the valve inlet, and a valve overflow leading out of the shut-off chamber. When the liquid level in the sample chamber rises to a predetermined level it disrupts the flow of liquid through the valve outlet and diverts later flow through the valve inlet into the shut-off chamber and through the valve overflow.

A further embodiment advantageously provides a first-void urine sampler which can be conveniently held by a user while they provide a urine specimen into a funnel, the sampler further comprising a valve which automatically directs a desired volume of first-void urine into a sample chamber and the remainder of the urine specimen to an exhaust passage for disposal.

The urine sampler may thus enable the user to urinate continuously without the need for bladder control or the potentially messy option of moving a collection receptacle out of the urine stream.

Advantageously, the valve provided within the sampler may seal the sample chamber after collection of the first portion of the liquid flow, or the first-void urine, until the sample chamber is removed from the sampler, even if the sampler is tilted or inverted with the sample chamber in place.

In preferred embodiments, the sampler may advantageously collect the first 2ml to 5ml or 10ml of a liquid flow or a urine specimen, although the parameters of the sampler may be varied in order to collect different liquid volumes. The sample collected may also be diluted to a limited degree by liquid or urine delivered later in the liquid flow or the urine specimen. This dilution depends on the speed of

operation and the effectiveness of the valve action of the sampler, and may depend on other parameters such as the rate of delivery of the liquid flow or the urine sample.

- 5 Clearly, the sampler may find applications in sampling the first portion of any liquid flow. The technique is not limited to urine sampling.

Description of Specific Embodiments and Best Mode

10

Specific embodiments of the invention will now be described by way of example, with reference to the accompanying drawings, in which:

- 15 Figure 1 is a three-quarter view of a urine sampler embodying the invention;

Figure 2 is a back view of the sampler of Figure 1;

- 20 Figure 3 is a left side view of the sampler of Figure 1;

Figure 4 is a front view of the sampler of Figure 1;

Figure 5 is a plan view of the sampler of Figure 1;

25

Figure 6 is a bottom view of the sampler of Figure 1;

Figure 7 is a vertical section of the sampler of Figure 1;

- 30 Figure 8 is a side view of a sampler substantially identical to that of Figure 1 showing the positions of transverse sections shown in Figures 9, 10 and 11; and

35 Figures 12 to 26 are vertical sections of a central portion of the sampler of Figure 1 in various stages of urine collection.

A preferred embodiment provides a urine sampler fabricated from plastic components, as shown in Figures 1 to 6 and in section in Figure 7. A substantially identical sampler, differing only in details of construction, is shown in figures 8 to 11. The samplers of these embodiments are fabricated from machined acrylic. For mass production, it may be preferable to make the sampler by injection moulding and to use a different plastics material suitable for moulding.

The dimensions of the sampler of the embodiment described below have been developed for the acrylic material. If the sampler is fabricated from a different material some modification of the sampler dimensions may be anticipated, particularly if the wetting properties of the material (for the liquid to be sampled) differ significantly from those of acrylic. The principles of operation of the sampler are materials independent, however, and only routine, non-inventive development within the capability of the person skilled in the art would be required to fabricate a sampler from a different material or for a different liquid in the light of the teaching herein.

The sampler 2 comprises an upper portion 4, a valve portion 6 and a sample chamber 8. The valve portion is a push fit onto the lower end of the upper portion, providing a liquid-tight seal, and is retained by a catch 10. The sample chamber is a push fit onto the valve portion, providing a liquid and air-tight seal. The catch ensures that the sample chamber can be removed without accidentally separating the upper and valve portions.

The sampler of the embodiment is designed for use in a substantially vertical orientation, with the sample chamber at the bottom. The following text will describe it in this orientation.

The upper end of the upper portion forms a funnel 12 for receiving urine. A handle 14 is provided for a user to hold the sampler while providing a urine specimen. Within the funnel, a baffle 16 is positioned to modulate the flow of urine and reduce turbulence at the funnel outlet 18. The handle, the baffle and a portion of the funnel may be moulded as a separate component and fastened onto the remainder of the upper portion by means of a clip 19 for convenience of manufacture.

The funnel outlet 18 drains into a first-catch reservoir 20 within the upper portion of the sampler. The first-catch reservoir is substantially cylindrical except at its lower end, where a frusto-conical tapered section 22 links it to a substantially cylindrical valve inlet 24, which is preferably of circular cross-section. When a urine specimen enters the funnel, it drains into the first-catch reservoir and flows through the valve inlet, in a cylindrical stream out of the upper portion of the sampler. The flow rate is determined primarily by the cross-section and lengths of the valve inlet and the tapered section and the head of liquid in the first-catch reservoir.

Below the level of the funnel outlet and at the upper end of the first-catch reservoir, a main overflow 26 leads to a vertical exhaust passage 28, which extends downwardly, substantially parallel to the first-catch reservoir and the valve inlet.

When the upper portion of the sampler is secured to the valve portion, the valve inlet forms the uppermost component of a valve 30. The structure of the valve can be seen in the sectional views of Figures 7, 9, 10 and 11.

At its lower end, the valve inlet leads into a shut-off chamber 32. This chamber is larger than the valve inlet in

horizontal cross-section and its side walls are set back from the walls of the valve inlet, so as not to interfere with the flow of urine out of the valve inlet. The shut-off chamber extends horizontally to a shut-off chamber overflow 34, which
5 drains into the exhaust 28.

A tapered passage 38 extends downwardly from the shut-off chamber, leading to a narrower, substantially cylindrical valve outlet 36 which opens into the sample chamber. The
10 valve outlet is aligned with the valve inlet and is preferably circular in section, having a slightly larger diameter than the valve inlet. Thus, in use, a urine stream from the valve inlet can initially pass through the shut-off chamber and the
15 valve outlet without touching the walls of either the shut-off chamber or the valve outlet.

The valve outlet is slightly offset to one side of the sample chamber. This allows space for a vent tube 40 which extends from within the sample chamber, upwardly through the shut-off
20 chamber and opens at its upper end 42 into a vent tube pocket 44. The lower end of the vent tube is blind but one or more vent holes 46 are defined in a side wall of the vent tube near its lower end. The holes preferably do not open either facing towards the centre of the sample chamber and the valve
25 outlet or in the opposite direction towards the closest portion of the sample chamber wall, but open laterally, into or out of the plane of the section in figure 7. This reduces the risk of liquid splashing in the sample chamber blocking the vent holes prematurely.

30 The vent tube is moulded as part of the valve portion of the sampler and, when the sampler is assembled, its upper end enters the vent tube pocket, which is a moulded recess in the upper portion of the sampler. When the sampler is assembled,
35 the vent tube pocket is connected to and extends upwardly from

the shut-off chamber, the upper end of the vent tube extending into the pocket alongside the valve inlet 24.

5 As shown in the transverse section of Figure 11, the vent tube need not be of circular section but is advantageously of an elongated or arcuate section to maximise the cross-sectional area of the vent while keeping it spaced from the valve outlet and the urine stream passing into the sample chamber. It is advantageous for the sample chamber to vent freely in order to
10 allow the first-void urine to flow as quickly as possible into the sample chamber. This reduces the risk of first-void urine mixing with later portions of the urine specimen in the first-catch reservoir.

15 Figures 12 to 26 illustrate the operation of the sampler of the embodiment during the collection of a first-void urine sample from a urine specimen. Each figure shows the sampler in cross-section, omitting its upper and lower ends in order to focus on the valve operation. Reference numerals
20 identifying components of the sampler are shown in Figure 12 but have been omitted from Figures 13 to 26 for clarity.

Figure 13 shows a urine specimen 100 entering the funnel and draining between the funnel and the baffle into the first void
25 chamber. Initially, the urine flows through the valve inlet 24 and passes as a cylindrical column, or jet, 102 into the sampler chamber. The diameter of the urine column, or jet, 102 is determined primarily by the shape and size of the valve inlet and is arranged so that the column passes directly
30 through the shut-off chamber and the valve outlet at this stage.

The flow rate through the valve inlet is predetermined to be slower than the flow rate into the funnel and therefore, as
35 shown in Figure 14, the level of liquid within the first void chamber 20, 22 above the valve inlet begins to rise as shown

in Figures 14 to 16, until it reaches a level 104 just below the main overflow 26, as shown in Figure 16. During this time, the liquid level 106 in the sample chamber rises. After this point, as shown in Figure 17, urine 108 begins to flow
5 into the main overflow and the exhaust 28, preventing the liquid level from rising further within the sampler.

At about this stage, as shown in Figure 17, the liquid level 110 in the sample chamber rises to cover the vent
10 holes 46 near the end of the vent tube 40. In Figure 17, this occurs at about the same time that the liquid in the first-catch reservoir overflows through the main overflow. This may not always be the case, however. For example, if the urine specimen is provided more rapidly, the first-catch
15 reservoir will overflow before the liquid level in the sample chamber covers the vent holes.

While the sample chamber is filling, up to the point at which the vent holes are covered, the air displaced from the sample
20 chamber by the liquid passes through the vent holes, along the vent tube, through the vent tube pocket and escapes through the shut-off chamber overflow. When the vent holes are covered, however, air can no longer escape along this route. The only escape route is then upwards through the valve outlet
25 (alongside the column of urine entering the sample chamber) and through the shut-off chamber overflow. As the air in the sample chamber flows up the valve outlet, against the
direction of flow of the urine, and the air pressure in the
sample chamber rises, the column of urine is disturbed and
30 bulges and touches the walls of the valve outlet. The initial bulging of the urine column is shown at 112 in Figure 17.

When the urine column wets the valve outlet around its entire circumference, no further air can escape from the sample
35 chamber. The combination of capillary attraction between the urine and the wall of the valve outlet and the increased air

pressure in the sample chamber prevents further flow of urine 114 through the valve outlet, as shown in Figure 18. This effectively closes the valve 30. As shown in Figures 18 to 20, no further urine enters the sample chamber and the shut-off chamber is filled by urine flowing through the valve inlet from the first-catch reservoir. This liquid covers the lower end of the vent tube pocket, providing a further seal to prevent air leaving the sample chamber. As shown in Figure 20, this liquid 116 escapes through the shut-off chamber overflow. At the same time, if urine is still entering the funnel, any excess 118 at the top of the catch reservoir escapes through the main overflow.

When delivery of the urine specimen into the funnel ceases, as shown in Figure 21 any liquid 120 above the main overflow drains through the main overflow while liquid in the first-catch reservoir and the shut-off chamber drains through the shut-off chamber overflow. This process continues as shown in Figures 22 to 25, until the first-catch reservoir and the shut-off chamber are empty. The final configuration of the sampler is shown in Figure 26, in which the desired sample is in the sample chamber and a small volume of urine 122 remains in the valve outlet, held by capillary attraction. The sample chamber may then be detached from the remainder of the sampler, without dislodging the liquid 122 within the valve outlet, for analysis or other purposes.

It can be seen that the urine sample in the sample chamber may advantageously contain a large proportion of first-void urine, subsequent parts of the specimen having been directed to the exhaust 28.

It can also be seen that in the final configuration of the sampler, shown in Figure 26, the sample is securely held within the sample chamber even if the sampler is tilted or inverted. Once the valve outlet is blocked by capillary

attraction and the vent holes are covered by the sample, the airlock within the sample chamber may advantageously prevent the sample from escaping either through the valve outlet or the vent tube.

5

Features and functions of the sampler

Funnel 12

10 The angle of the funnel is selected to encourage efficient flow into the first-catch reservoir. If the funnel angle is too shallow, undesirable mixing of urine in the funnel may occur. However, if the funnel angle is too steep, urine may be forced too quickly into the first-catch reservoir, causing
15 turbulence and mixing in the reservoir.

In the embodiment, the funnel angle, between two opposite walls of the funnel, is 40 degrees.

20 Baffle 16

The baffle should prevent urine jetting directly into the first-catch reservoir, which would cause mixing, or into the overflow, which would risk losing the first-void urine. The
25 profile of the baffle should be selected to encourage tangential flow, rather than vertical flow, within the sampler, to reduce mixing and encourage flow into the main overflow when the first-catch reservoir has filled. This
30 assists in achieving the object of the sampler, of collecting the first few millilitres of urine in the specimen.

When a urine specimen first enters the sampler, it enters the first-catch reservoir and immediately begins to flow into the sample chamber. The valve inlet restricts the urine flow rate
35 and so, as the sample chamber fills, the urine level in the first-catch reservoir rises. It is therefore desirable to

limit mixing of urine in the first-catch reservoir, otherwise later portions of the urine specimen might mix with first-void urine at the lower end of the first-catch reservoir and enter the sample chamber, disadvantageously diluting the sample. The design of the baffle can help to reduce such mixing by encouraging horizontal, rather than vertical, flow directions within the sampler and especially within the first-catch reservoir. Advantageously, limiting vertical flow can lead to layering of the urine in the first-catch reservoir, with more concentrated first-void urine at the bottom, which will flow into the sample chamber, and urine containing a higher proportion of later parts of the urine specimen at the top.

In the embodiment, the clearance between the baffle end and the inside of the funnel should be between 1.5mm and 3.0mm. Clearance less than 1.5mm tends to restrict flow and lead to urine collecting above the baffle, which disrupts flow into the sampler. Clearance more than 3mm tends to lead to a risk of the urine specimen jetting directly into the main overflow or the first-catch reservoir.

First-catch reservoir

The reservoir traps the first-void urine however quickly the specimen is delivered and limits mixing with later portions of the specimen, which escape through the main overflow. In practice, about the first 1ml or 2ml of the sample enters the sample chamber rapidly, while the first-catch reservoir fills. If the desired sample size is, say, 4 ml, the balance of the sample is contained at that stage in the lower part of the first-catch reservoir, and drains progressively into the sample chamber. Figures 13 to 16 illustrate this process in the embodiment. When the first-catch reservoir fills, its upper part contains liquid which is not intended to form part of the sample but which effectively forms a buffer to protect the liquid in the lower part of the first-catch reservoir from

turbulence and mixing caused by further urine entering the sampler, and therefore helps to reduce dilution of the sample. This is a facet of the phenomenon of layering discussed above. Thus, the capacity of the first-catch reservoir should
5 advantageously be greater than the desired sample volume minus the volume of the part of the sample which enters the sample chamber before the first-catch reservoir has filled plus the volume of the buffer liquid. In effect, for sample volumes of a few millilitres this means that the capacity of the
10 first-catch reservoir advantageously approximately equals the desired sample volume to be collected.

The diameter of the first-catch reservoir should be selected to prevent or reduce mixing between urine at the bottom of the
15 reservoir and urine higher in the reservoir, which may occur if the diameter is too large. (The issue of mixing and layering in the first-catch reservoir is also discussed above in relation to baffle design). However, if the first-catch reservoir diameter or volume is too small, then a portion of
20 the first-void urine may be lost through the main overflow.

As described above the first-catch reservoir preferably has a minimum volume to prevent dilution of the sample, but it is also desirable that its diameter or cross-section is not too
25 large. There may therefore need to be a compromise between these dimensions in order to prevent the height of the first-catch reservoir, and therefore the height of the sampler, being undesirably large.

30 In the embodiment, the diameter of the cylindrical portion of the reservoir is 10mm and its height is 50mm. This is appropriate for a 4ml sample volume. For different sample volumes of a few millilitres, the height or other dimensions of the first-catch reservoir may advantageously be varied so
35 that the reservoir volume is about the same as the sample volume.

First-catch reservoir, frusto-conical section 22

This lower section of the first-catch reservoir, which may be termed the reservoir funnel, blends the diameter of the first-catch reservoir down to the diameter of the valve inlet. The angle of the reservoir funnel should be selected as follows. If the angle is too shallow, flow into the valve inlet may be inhibited. If the angle is too steep, it may encourage excessively rapid flow through the valve inlet. In the embodiment, the reservoir funnel is 10mm in length, blending from the 10mm diameter of the first-catch reservoir to the 3.2mm diameter of the valve inlet.

Valve inlet

In the embodiment, this is a circular cylinder of 3.2 millimetre diameter. This provides a focussed jet, or column, of urine, through the valve outlet into the sample chamber. The diameter of the valve inlet should not be too small, in which case the jet of urine it produces may puncture through the drop of urine in the valve outlet, which is held by capillary action and prevents further flow into the sample chamber after the sample has been collected.

Tapered passage 38 leading to the valve outlet

As described above, the urine column from the valve inlet initially passes through the valve outlet directly into the sample chamber. At this stage it is important that the urine flows freely into the sample chamber and so it is desirable to prevent the urine column from wetting the entire circumference of the valve outlet, which might cause formation of a plug of liquid held by capillary action in the valve outlet and prematurely disrupt or prevent further urine flow into the sample chamber.

If the sampler is tilted during collection of a specimen, the urine column produced by the valve inlet may curve downwards. The provision of a tapered entrance to the valve outlet may increase the tolerance of the sampler to being tilted during use by reducing the chance or the extent of contact between the urine column and the circumference of the valve outlet or the floor of the shut-off chamber beside the valve outlet.

Valve outlet 36

The dimensions of the valve outlet may advantageously be selected in view of the following factors. Its diameter should be small enough to enable the collected sample in the sample chamber to remain sealed by the surface tension in the suspended urine drop, or plug, within the valve outlet. The tapered passage 38 leading into the valve outlet helps to cope with any irregularities in the urine stream exiting the valve inlet, which may arise due to turbulence or tilting of the sampler.

A further factor is the relationship between the sizes and positions of the valve inlet and the valve outlet. In the embodiments described above the valve outlet is aligned with and of slightly larger diameter than the valve inlet. This is one way to achieve the functional requirements of this aspect of the sampler that the urine column from the valve inlet should pass freely through the valve outlet until, at a

predetermined liquid level in the sample chamber, the column is disrupted and a plug of liquid forms in the valve outlet, causing further urine to be diverted into the shut-off chamber. Thus, the valve outlet may be of any size or orientation as long as these functional requirements are met; it may, for example, be tilted relative to the valve inlet so that the urine initially strikes and flows along the wall on one side of the valve outlet into the sample chamber.

In the embodiment, the valve outlet is a circular cylinder of 4.2mm diameter and 8.5mm long. The tapered passage at its entrance is 3.5mm long and flares at an angle of 30 degrees.

5 Vent holes 46

Each vent hole should be sufficiently small that when the urine level in the sample chamber rises to cover the vent holes, the urine is prevented by surface tension from entering
10 the vent tube. The cumulative cross-section of all of the vent holes (or slots) should be large enough to allow air to escape through the vent tube fast enough to allow the sample chamber to fill; it is desirable to allow the sample chamber to fill sufficiently quickly to prevent unnecessary mixing of
15 the first-void urine in the first-catch reservoir with later portions of the urine specimen.

It may be advantageous to use slot-shaped vent holes, rather than circular vent-holes for example, to increase air flow
20 while reducing the minimum lateral dimension of the vent-holes and therefore making it easier for surface tension to block the vent-holes.

In the embodiment, the maximum dimension for any one of the
25 vent holes is 0.5mm, to prevent urine ingress into the vent tube, and the minimum total open area of all of the vent holes is 0.8mm², to provide sufficient air flow rate.

Vent tube 40

30 As described above, the lower end of the vent tube in the embodiment may be sealed and vent holes advantageously defined in the side of the vent tube near its end and on the side of the vent tube facing neither the urine column entering the
35 sample chamber nor the wall of the sample chamber furthest from the entering urine column. When the urine enters the sample chamber, it tends to splash, rising particularly high

adjacent the wall of the sample tube furthest from the valve outlet. This is driven by the tendency of the urine stream to rotate at the bottom of the sample tube, following the wall of the sample chamber. The positioning of the vent holes on the sides of the vent tube therefore helps to control unwanted or premature splashing of the vent holes.

The upper end of the vent tube nests into the vent tube pocket. It should be noted that when the shut-off chamber floods after collection of the sample in the sample chamber, the lower end of the pocket is sealed by the presence of the urine, which helps to maintain the air pressure within the sample chamber, which in turn prevents further access of urine into the sample chamber.

In the embodiment, the vent pocket wall is spaced from the vent tube by between 0.25mm and 0.5mm.

Shut-off chamber 32

The height of the shut-off chamber determines the distance between the valve inlet and the valve outlet, and so may affect the relationship between their diameters. The capacity of the shut-off chamber also affects how quickly and effectively the vent pocket is sealed after the sample is collected. Further, the height of the shut-off chamber affects the tolerance of the sampler to tilting during use, given that the urine stream leaving the valve inlet must pass through the shut-off chamber and the valve outlet as described above.

If the shut-off chamber volume is too large or its height is too great, then after urine stops flowing into the sample chamber and starts flowing into the shut-off chamber, it may take too long to reach and cover the vent-tube pocket. This may lead to unreliability as, until the vent-tube pocket is covered, the sealing of the sample chamber depends on the plug

of liquid in the valve outlet and the capillary action sealing the vent holes. If the shut-off chamber volume is too small or its height too small, then if the urine column flowing from the valve inlet accidentally hits the edge of the valve outlet within the shut-off chamber, sufficient liquid may collect in the shut-off chamber to cover the vent-tube pocket and cause the valve to close prematurely. Again, this is a reliability issue, which may lead to inconsistent operation if, for example, the sampler is tilted or shaken excessively during urine collection.

It will be appreciated that other features such as the tapered entrance passage of the valve outlet may also affect these aspects of the sampler's operation; the shape and size of the tapered passage influence the risk of the urine column hitting the floor of the shut-off chamber adjacent the edge of the valve outlet (or its tapered entrance passage) and prematurely deflecting liquid into the shut-off chamber.

In the embodiment, the shut-off chamber is 3.5mm high and has a volume of approximately 320mm³.

Shut-off chamber overflow 34

This overflow allows urine from the shut-off chamber to escape to the exhaust passage. The cross-section of the overflow must not be too large, otherwise it will empty the shut-off chamber too quickly, which could break the seal provided by the urine in the shut-off chamber at the base of the vent pocket. If the shut-off chamber overflow is too small, it may not allow sufficient flow through the shut-off chamber to block the vent pocket quickly enough when the sample chamber is filled to the required level.

In the embodiment, the shut-off chamber overflow is a circular cylindrical passage of diameter 2.4mm.

Sampler characteristics

The sampler of the embodiment is designed to collect a 2ml sample. It is designed to operate at urine flow rates between 15ml and 30ml per second. For other applications, the sizes of the components of the sampler may be altered appropriately, for example to collect samples of 5ml or 10ml. It is desirable that the sample should be as pure as possible, ideally containing only the desired volume of first-void urine. Usually, however, some dilution of the sample with later portions of the urine specimen occurs due to mixing in the first-catch reservoir. Tests have, for example, achieved collection of 10ml samples in which each sample was collected from amongst the first 12ml of the (much larger) urine specimen.

For reliable operation, it is important that the internal surfaces of the sampler are clean and dry because wetting of components such as the valve outlet and the shut-off chamber should be predictable. In one embodiment, the sampler may be manufactured as a disposable item and kept in a sealed container before use, ensuring that it is clean and dry when used. If a sampler were to be reused, it may be important to be able to dismantle it for effective cleaning and drying.

Further Embodiments

In a simpler embodiment, the sample chamber is not provided with a vent-tube extending into the chamber, while the valve outlet is lengthened by a tubular portion extending into the sample chamber. While the sample chamber is filling, the liquid column from the valve inlet passes through the valve outlet and air from the chamber vents through the valve outlet alongside the liquid column. When the urine level in the sample chamber reaches the end of the urine valve outlet, no further air can escape and the liquid column in the valve outlet is disrupted, wetting the circumference of the valve

outlet and forming a liquid plug. This closes the valve, diverting the urine flowing from the valve inlet into the shut-off chamber and through the shut-off chamber overflow.

Claims

1. A sampler for sampling a first portion of a liquid flow, in which the first portion passes through a valve into a sample chamber, the valve comprising;
 - 5 a valve inlet coupled to an inlet of the sampler;
 - a valve outlet opening into the sample chamber, the valve outlet being positioned, in use, substantially below the valve inlet;
 - a shut-off chamber separating the valve inlet from the
 - 10 valve outlet; and
 - a valve overflow leading out of the shut-off chamber.
2. A sampler according to claim 1, in which the valve outlet is substantially the same shape, in transverse section, as the valve inlet, and is of larger transverse dimensions.
- 15 3. A sampler according to claim 1 or 2, in which the valve outlet is aligned with the valve inlet.
4. A sampler according to any preceding claim and further comprising;

- 20 a vent tube extending between a vent inlet and a vent outlet, the vent inlet being positioned within the sample chamber, and the vent outlet being positioned, in use, at a level above the valve outlet.
- 5. A sampler according to claim 4, in which the level of the vent inlet within the sample chamber determines the level
- 25 of liquid to be collected within the sample chamber in that, in use, the valve closes when liquid in the sample chamber

reaches the vent inlet and obstructs it, substantially preventing further venting of air from the sample chamber.

6. A sampler according to claim 4 or 5, in which the vent outlet opens into a vent-tube pocket which extends, in use,
5 upwardly from the shut-off chamber.

7. A sampler according to any preceding claim, in which, after collection of the sample in the sample chamber, further liquid entering the valve during use drains through the valve overflow.

10 8. A sampler according to any preceding claim, in which the sampler inlet comprises a first-catch reservoir opening into the valve inlet.

9. A sampler according to claim 8, in which the sampler inlet comprises a main overflow positioned, during use, at an
15 upper end of the first-catch reservoir.

10. A sampler according to claim 8 or 9, in which the sampler inlet comprises an entrance funnel opening into the first-catch reservoir, the funnel optionally being provided with a baffle for reducing turbulence within the first-catch
20 reservoir.

11. A sampler according to claim 9, in which the shut-off chamber overflow and the main overflow lead into a common overflow outlet of the sampler.

12. A sampler according to any preceding claim, in which the
25 sample chamber is removable.

13. A method for sampling a first portion of a liquid flow using a sampler as defined in any preceding claim, comprising the steps of;

operating the sampler so that liquid initially flows through the valve inlet and then passes through the shut-off chamber and the valve outlet;

collecting liquid in the sample chamber until it reaches
5 a predetermined level at which the flow of liquid through the valve outlet is disrupted so that it touches and wets the wall of the valve outlet, blocks the valve outlet and shuts the valve, preventing further flow of liquid into the sample chamber; and

10 diverting further liquid flowing through the valve inlet to flow through the shut-off chamber and the valve overflow.

14. A method according to claim 13, in which the predetermined level is at an exit of the valve outlet, the liquid flow being disrupted by the liquid in the sample
15 chamber covering the exit of the valve outlet.

15. A method according to claim 13 or 14, using a sampler which further comprises a first-catch reservoir leading into the valve inlet and a main overflow positioned, in use, at an upper end of the first-catch reservoir, comprising the steps
20 of;

passing liquid into the first-catch reservoir, which then flows into the valve inlet;

draining excess liquid through the main overflow when the first-catch reservoir is full; and

25 after the valve shuts, draining the first-catch reservoir through the valve inlet, the shut-off chamber and the valve overflow.

16. A valve for a liquid sampler as defined in any of claims 1 to 12.

17. A liquid sampler substantially as described herein, with reference to the drawings.

5 18. A valve substantially as described herein, with reference to the drawings.

19. A method for sampling liquid substantially as described herein, with reference to the drawings.

20. A first-void urine sampler comprising;

10 a sample chamber for collecting a first-void urine sample; and

a valve for allowing the sample to enter the sample chamber before the valve closes to divert later-voided urine away from the sample chamber.

15 21. A method for diagnosis comprising the steps of; collecting a first-void urine sample; and

testing the sample.

ABSTRACT

Liquid Sampler and Method (figure 7)

In a sampler for sampling a first portion of a liquid flow,
the liquid flow enters at a sampler inlet (12). The sample
5 passes through a valve (30) into a sample chamber (8) and then
the valve closes, diverting the remainder of the liquid flow
to a sampler overflow (6). The valve comprises a valve inlet
(24) coupled to the sampler inlet, a valve outlet (38, 36)
opening into the sample chamber and positioned, in use,
10 substantially below the valve inlet, a shut-off chamber (32)
separating the valve outlet from the valve inlet, and a valve
overflow (34) leading from the shut-off chamber to the sampler
overflow (6). When the liquid level in the sample chamber
rises to a predetermined level it disrupts the flow of liquid
15 through the valve outlet and diverts later flow through the
valve inlet into the shut-off chamber and through the valve
overflow.

1/12

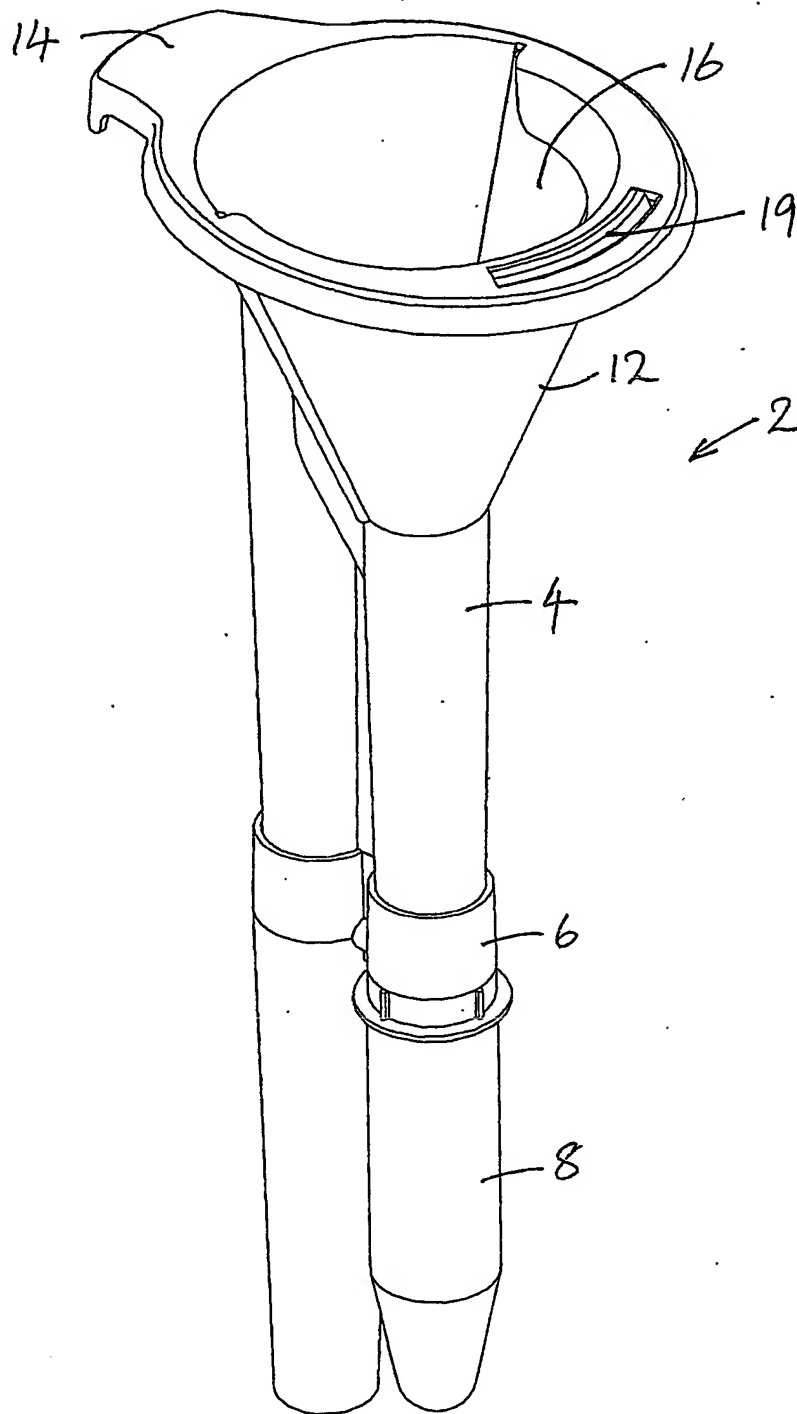


FIGURE 1

2/12

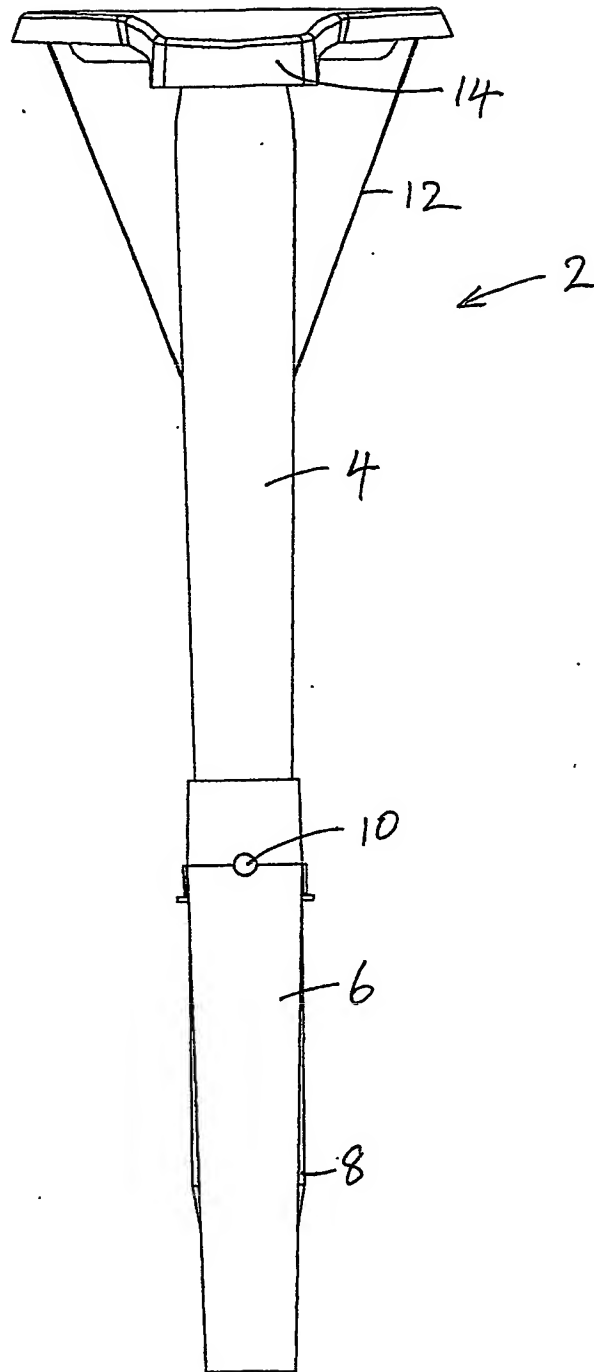


FIGURE 2

3/12

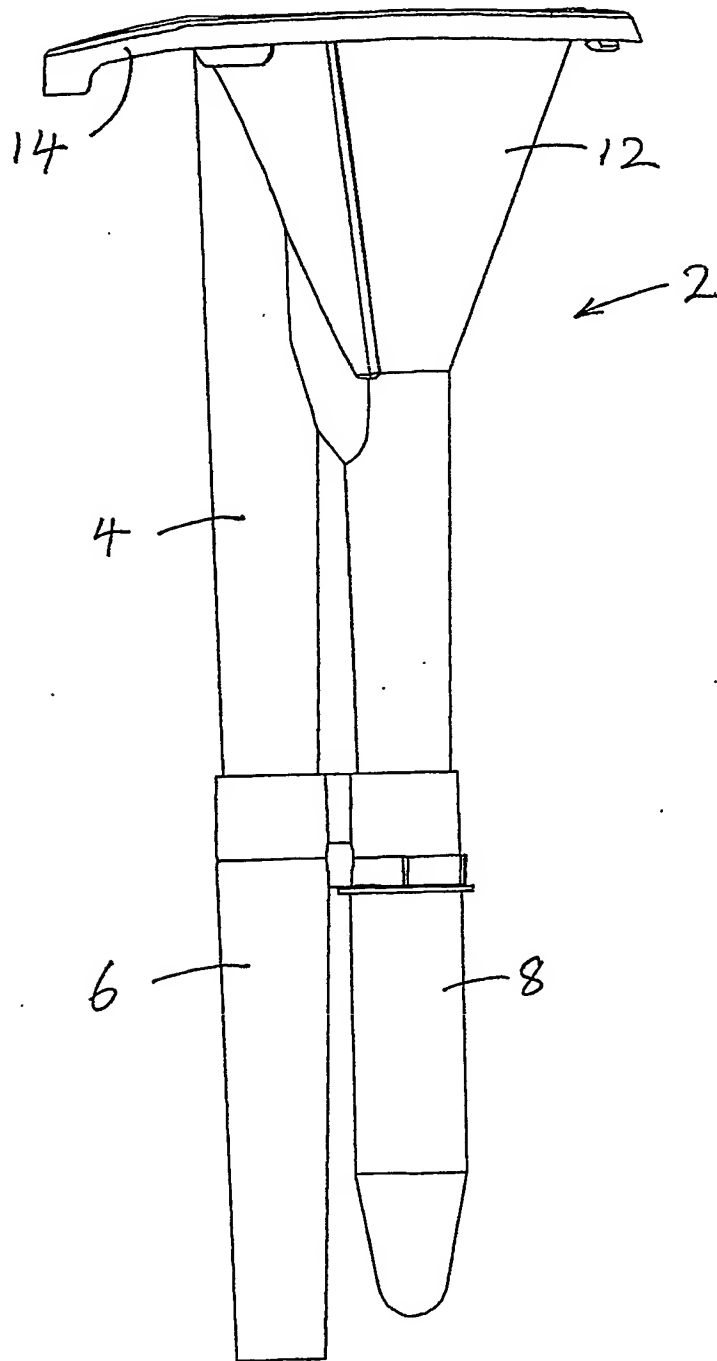


FIGURE 3

4/12

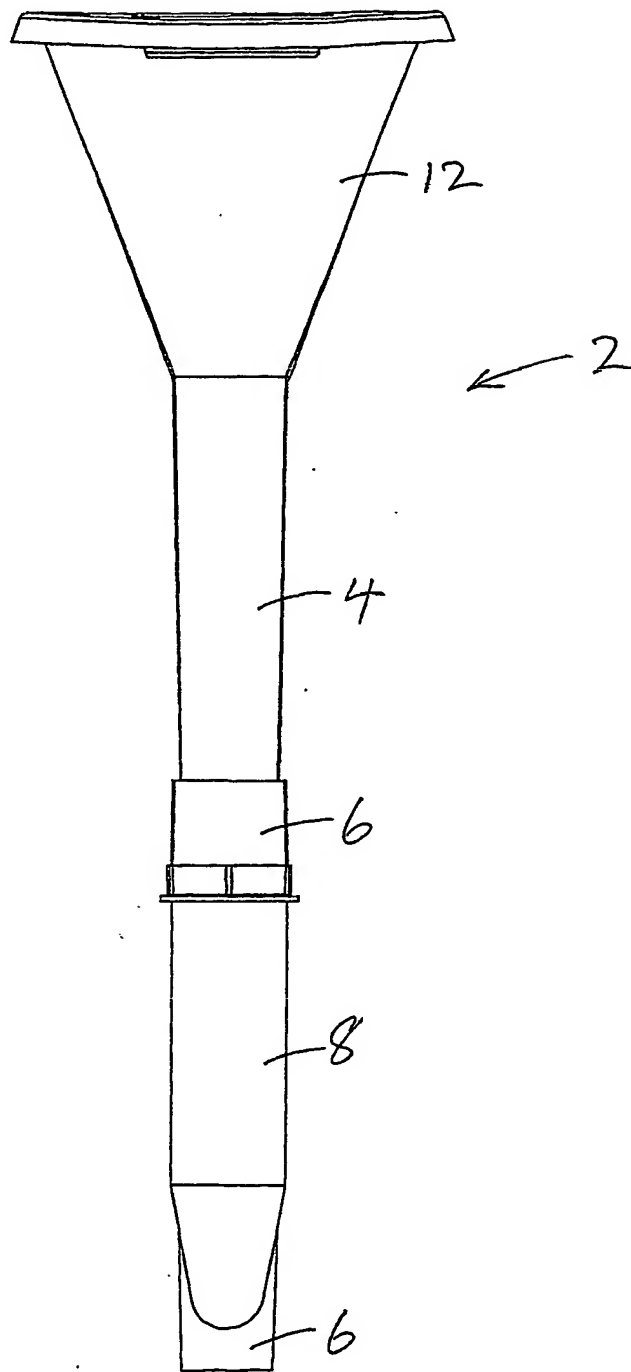
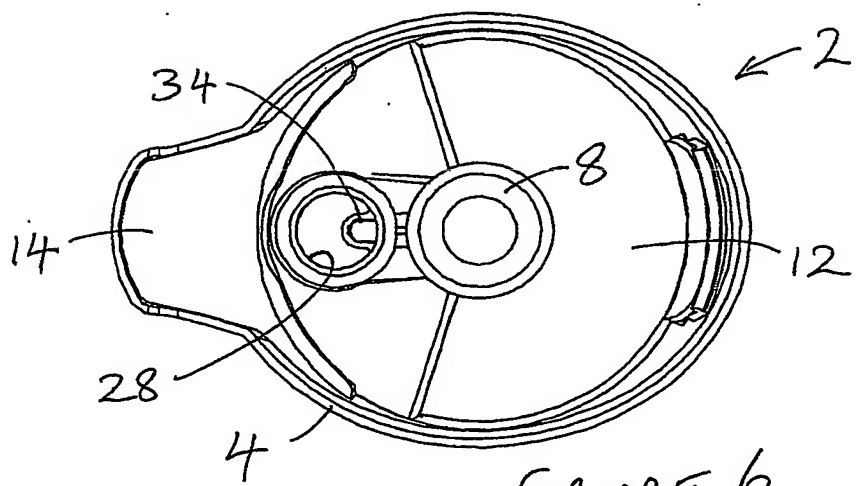
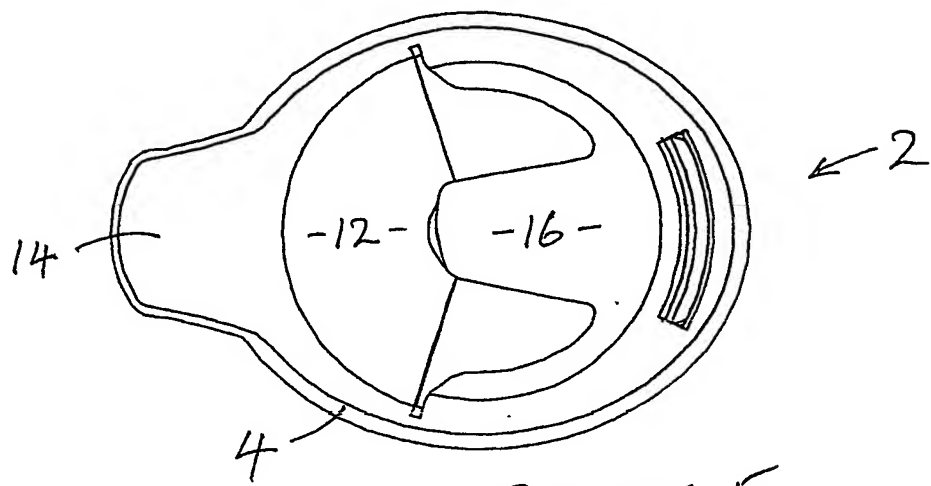


FIGURE 4

5/12



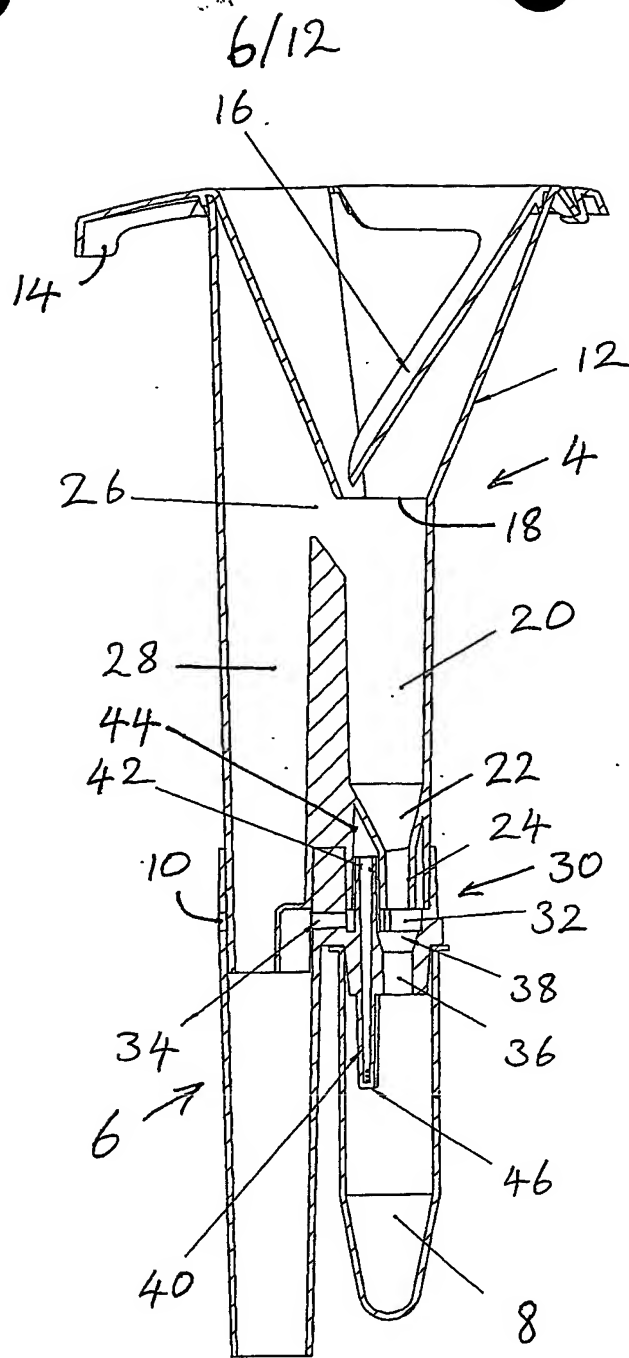
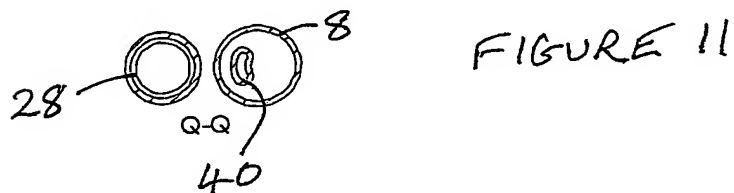
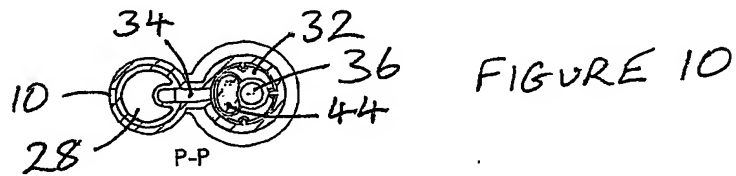
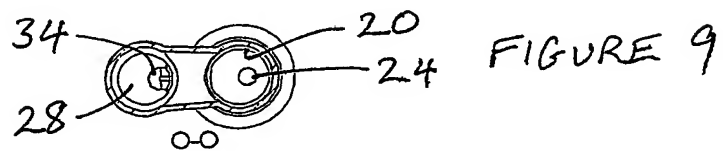
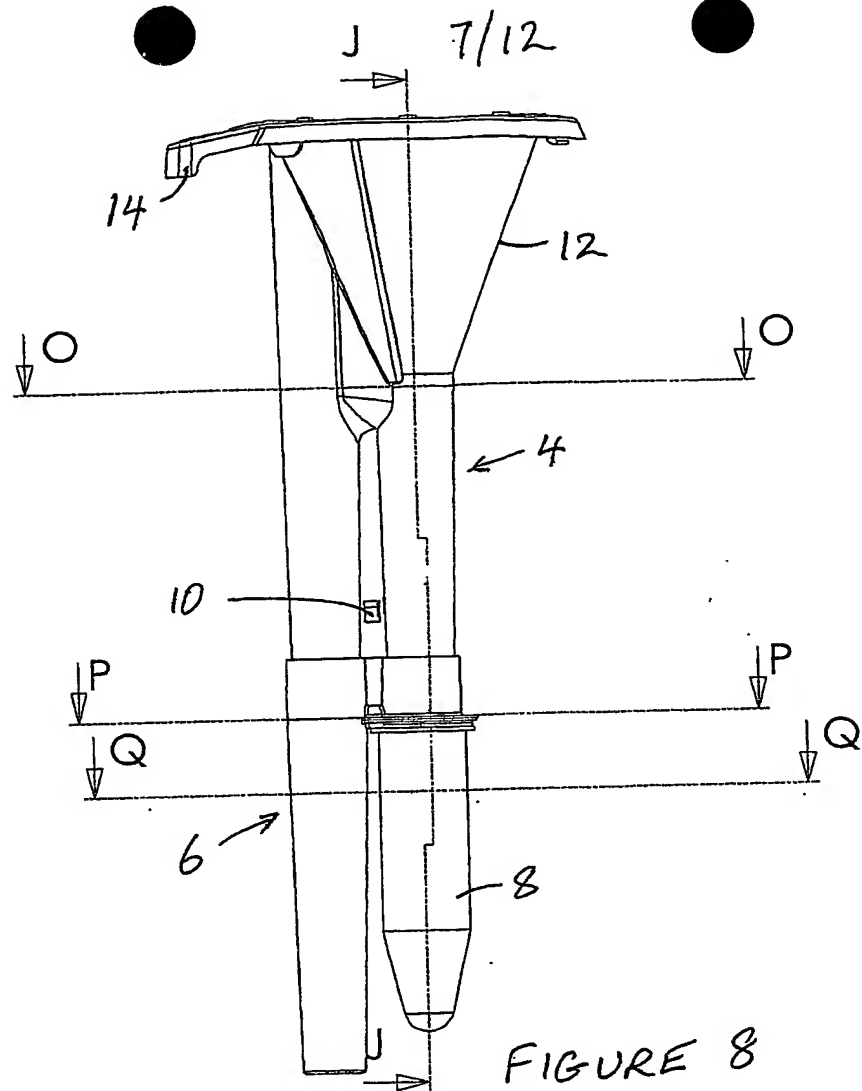


FIGURE 7



8/12

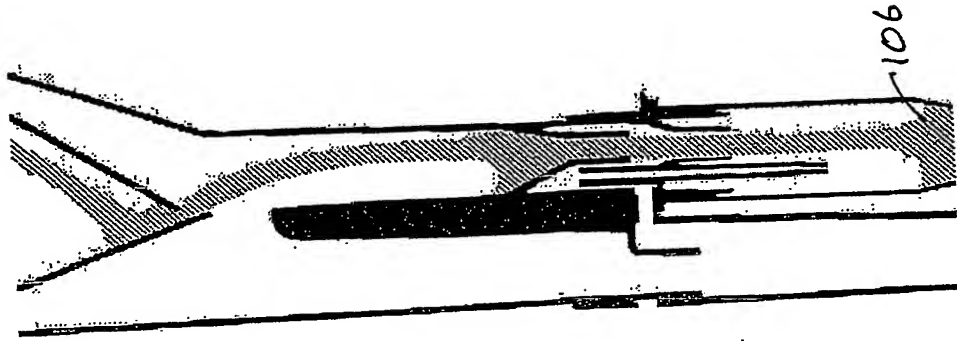


FIGURE 14

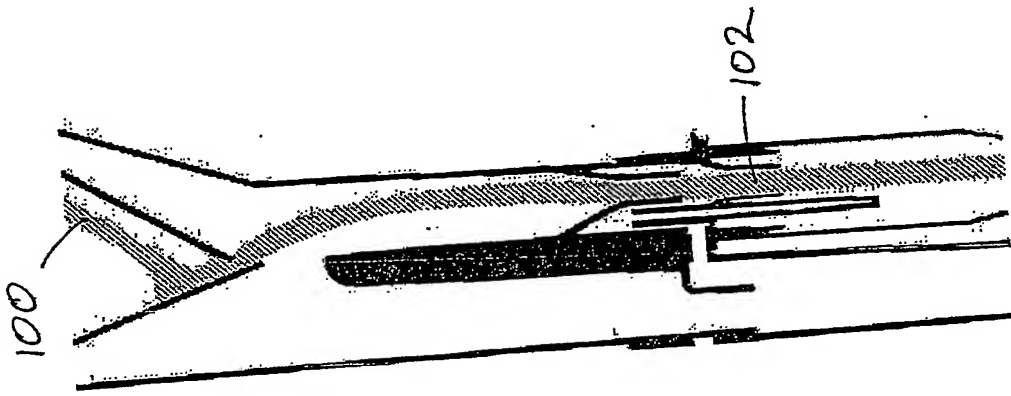


FIGURE 13

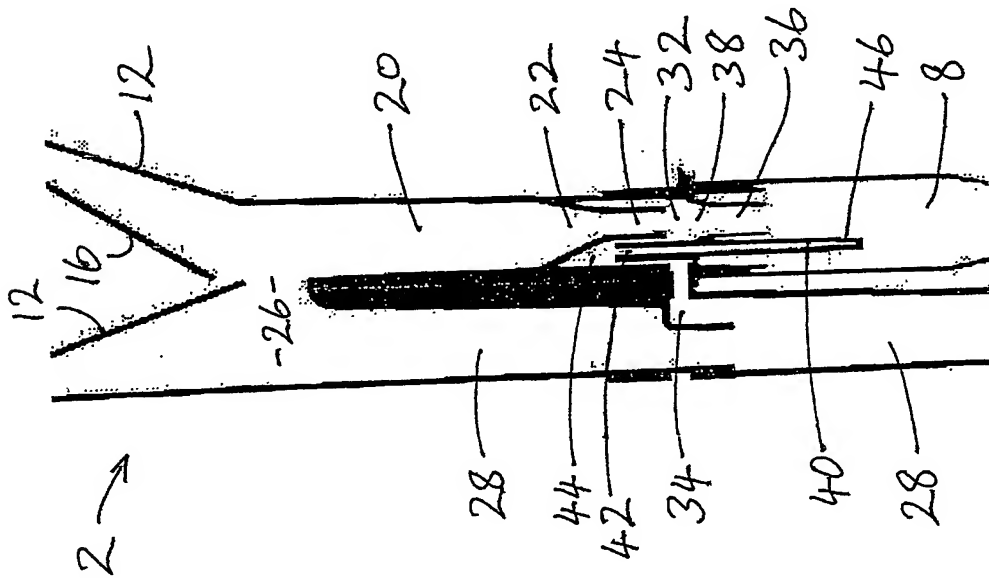


FIGURE 12

9/12

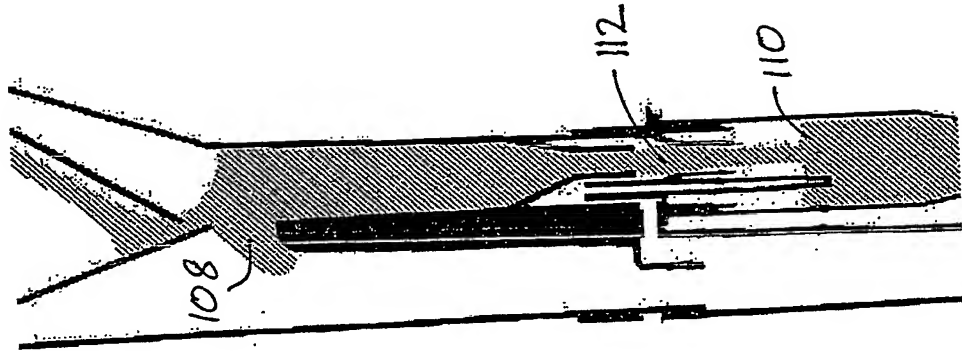


FIGURE 17

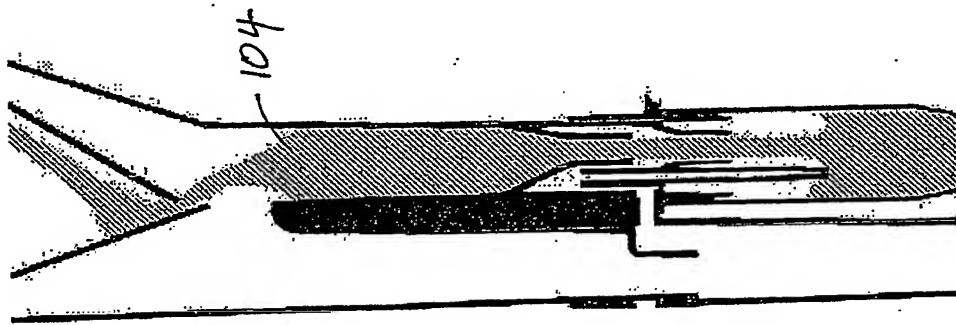


FIGURE 16

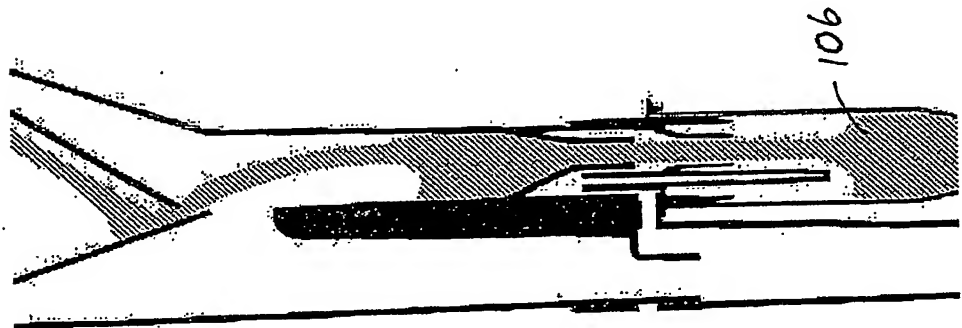


FIGURE 15

10/12

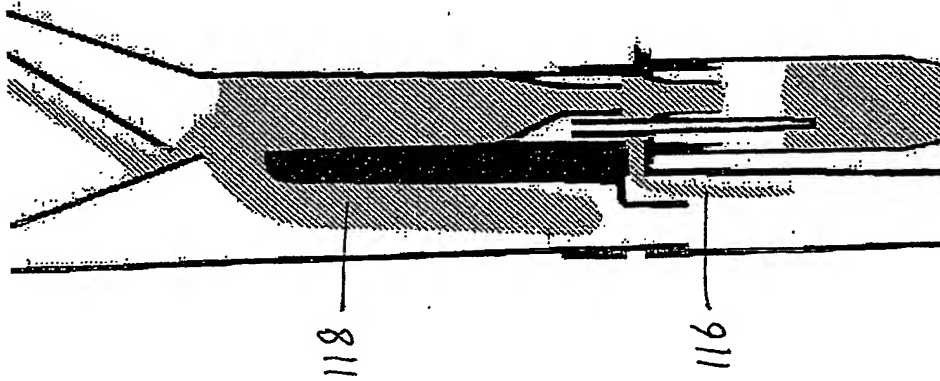


FIGURE 20

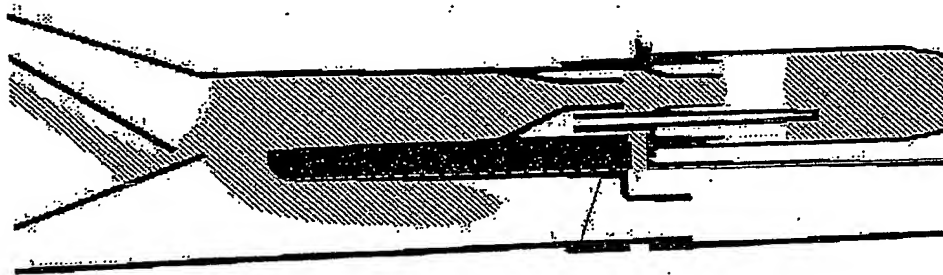


FIGURE 19

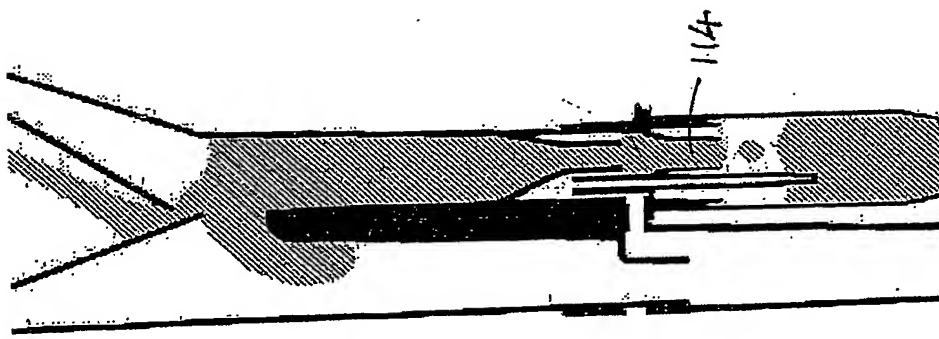


FIGURE 18

11/12

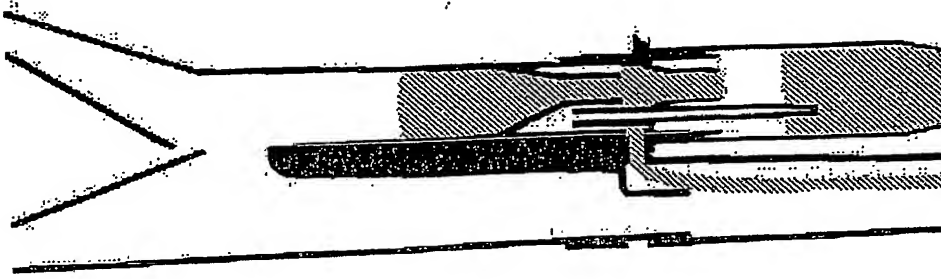


FIGURE 23

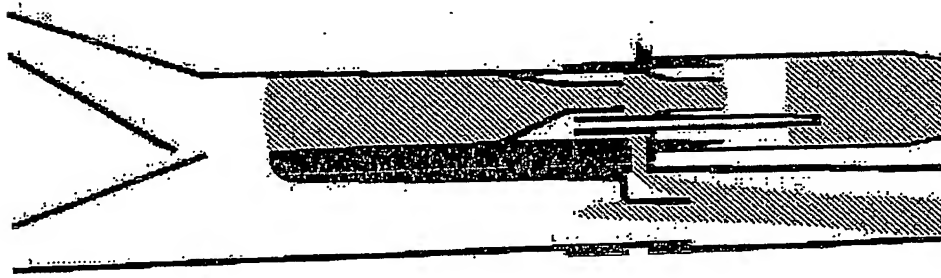


FIGURE 22

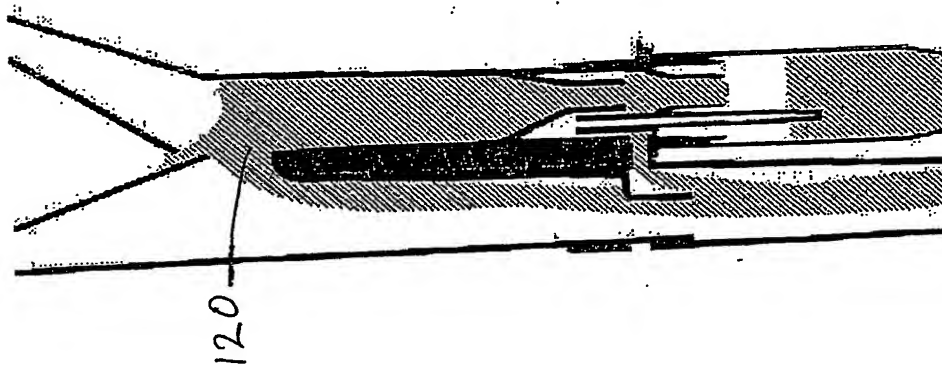


FIGURE 21

12/12

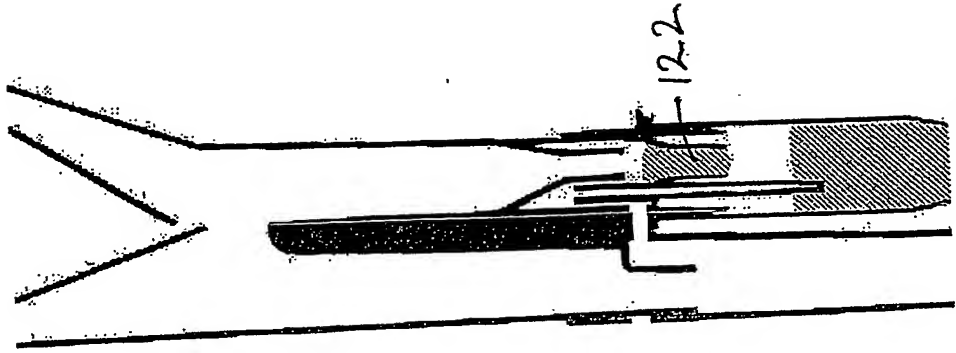


FIGURE 26

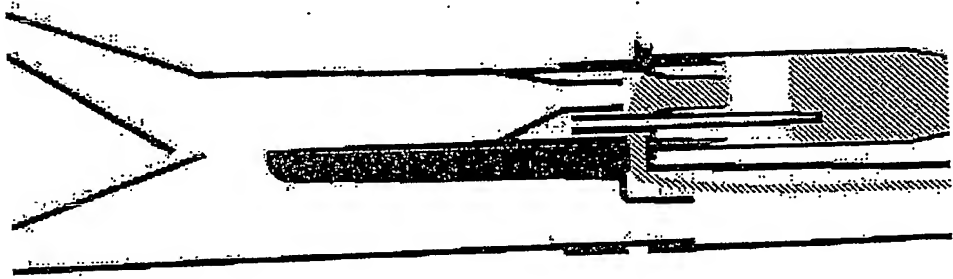


FIGURE 25

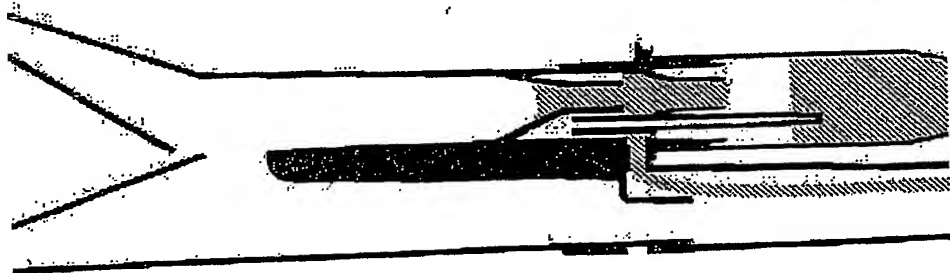


FIGURE 24

**This Page is Inserted by IFW Indexing and Scanning
Operations and is not part of the Official Record**

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

☐ **BLACK BORDERS**

☐ **IMAGE CUT OFF AT TOP, BOTTOM OR SIDES**

☐ **FADED TEXT OR DRAWING**

☐ **BLURRED OR ILLEGIBLE TEXT OR DRAWING**

☐ **SKEWED/SLANTED IMAGES**

☐ **COLOR OR BLACK AND WHITE PHOTOGRAPHS**

☐ **GRAY SCALE DOCUMENTS**

☒ **LINES OR MARKS ON ORIGINAL DOCUMENT**

☐ **REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY**

☐ **OTHER:** _____

IMAGES ARE BEST AVAILABLE COPY.

As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.